## **AMENDMENTS TO THE CLAIMS:**

This listing of claims will replace all prior versions and listings of claims in the application:

## **LISTING OF CLAIMS:**

- 1. (currently amended): A protein comprising <u>a</u> the sequence <u>selected from the group consisting of Seq ID No 1 or Seq ID No 4, or a variant thereof, SEQ ID NO:1, a variant of SEQ ID NO:1, SEQ ID NO:4, and a variant of SEQ ID NO:4, wherein the <u>sequence is</u> capable of hydrolysing sphingomyelin.</u>
- 2. (currently amended): The protein according to claim 1, wherein the Seq ID No-1 or Seq ID No-4 or variant thereof sequence is capable of hydrolysing sphingomyelin at pH 7.5-9.
- 3. (currently amended): The protein according to claim 1, wherein the  $\frac{\text{Seq ID No 1}}{\text{or Seq ID No 4 or variant thereof sequence}}$  has  $\frac{\text{Seq ID No 4 or variant thereof sequence}}{\text{sequence}}$  has  $\frac{\text{Seq ID No 4 or variant thereof sequence}}{\text{sequence}}$  has  $\frac{\text{Seq ID No 4 or variant thereof sequence}}{\text{sequence}}$  has  $\frac{\text{Seq ID No 4 or variant thereof sequence}}{\text{sequence}}$  has  $\frac{\text{Seq ID No 4 or variant thereof sequence}}{\text{seq ID No 4 or variant thereof sequence}}$  of its hydrolysing activity at pH  $\frac{\text{Seq ID No 4 or variant thereof sequence}}{\text{seq ID No 4 or variant thereof sequence}}$ .
- 4. (currently amended): The protein according to claim 1, wherein the variant of SEQ ID NO:1 has at least 80% identity with SEQ ID NO:1 and the variant of SEQ ID NO:4 has at least 80% identity with SEQ ID NO:4 Seq ID No 1 or Seq ID No SEQ ID NO: 4.
- 5. (currently amended): A nucleotide sequence encoding the protein according to claim 1 any of claims 1-4.
- 6. (currently amended): The nucleotide sequence according to claim 5, wherein the nucleotide sequence comprises Seq ID No SEQ ID NO: 2 or Seq ID No SEQ ID NO: 5.
- 7. (currently amended): A recombinant expression and secretion vector, comprising: a polynucleotide encoding a secretion signal peptide;

- a DNA sequence which promotes transcription in a host cell located upstream from the polynucleotide encoding the secretion signal peptide;
- a DNA sequence encoding a protein according to <u>claim 1</u> any of claims 1-4 in a translation reading frame with said polynucleotide encoding the secretion signal peptide; and
- a transcription terminator sequence located downstream from the DNA sequence encoding said protein.
- 8. (currently amended): A host cell, comprising: the recombinant expression system according to claim 7, wherein the host cell expresses from which Alk-Smase is expressed.
- 9. (currently amended): The host cell according to claim 8, wherein the host cell is selected from the group consisting of a bacteria, a mammalian cell or and a yeast cell; and which in the absence of the recombinant expression system according to claim 7, the host cell does not normally produce an Alk- Smase.
- 10. (currently amended): A method for isolation of human Alk-Smase protein, the method comprising the steps of:
- xvi) providing a small intestinal or colon content from a human;;
- xvii) homogenising the small intestinal or colon content;
- xviii) purifying Alk-Smase from the homogenized content using DEAE Sephadex chromatography;
- xix) purifying the Alk-Smase using Uno anion exchange chromatography; and xx) purifying the Alk.Smase using hydrophobic chromatography, thereby isolating the human Alk-Smase protein.
- 11. (currently amended): A method for preparation of recombinant Alk-Smase protein capable of hydrolysing sphingomyelin, the method comprising the steps of :

  ix) providing a host cell according to claim 8 any of claims 8-9 and a host cell growth medium; ;
- x) preparing a host cell culture;

- xi) culturing the host cell culture; and xii) harvesting the host cell culture and recovering the human recombinant Alk- Smase.
- 12. (currently amended): The method according to claim 11, wherein the Alk-Smase protein is recovered either from the culture medium, or the host cells-or after separating the host cells from the culture medium.
- 13. (currently amended): An isolated Alk-Smase protein, comprising the protein according to claim 1 any of claims 1-4, having, wherein the protein has an active site with the amino acid sequence AFVTMTSPCHFTLVTGKY (Seq ID No SEQ ID NO: 3) or a variant thereof.
- 14. (currently amended): A composition, comprising: a protein according to claim 1; any of claims 1-4, or a nucleic acid according to any of claims 5-6, or an isolated Alk-Smase according to any of claims 12-13, and a biocompatible carrier or additive.
- 15. (currently amended): Use of A method for treating colon cancer, comprising: administering a composition comprising at least one of a protein according to claim 4, any of claims 1-4, or a nucleic acid according to claim 5, and any of claims 5-6, or an isolated Alk-Smase according to claim 12 to a patient any of claims 12-13, for the preparation of a pharmaceutical composition for the treatment of colon cancer.
- 16. (currently amended): A kit comprising:
  the protein according to claim 1 any of claims 1-4, or the isolated protein according to claim 13; and
  a stabiliser.
- 17. (original): The kit according to claim 16, wherein the protein is in a lyophilised form or freeze-dried form.
- 18. (new): The method according to claim 12, wherein the Alk-Smase protein is recovered after separating the host cells from the culture medium.

- 19. (new): A composition, comprising: a nucleic acid according to claim 5; and a biocompatible carrier or additive.
- 20. (new): A composition, comprising: an isolated Alk-Smase according to claim 12; and a biocompatible carrier or additive.